IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Examiner : Kantamneni, Shobha

Art Unit : 1617

Applicants : Kivlighn et al.
Serial No. : 09/892,505
Filed : June 28, 2001

For : Treatment For Cardiovascular Disease

DECLARATION OF BERNARDO RODRIGUEZ-ITURBE, M.D.

I, Bernardo Rodriguez M.D., hereby declare and say as follows:

THAT, I am Professor of Medicine at Zulia University in Maracaibo, Venezuela, and President-Elect for the International Society of Nephrology. I am a world expert in the field of high blood pressure and nephrology (see my curriculum vitae attached) and have special expertise in animal models of hypertension. I am well versed with the work of Dr Johnson as I have collaborated with him on a variety of projects.

THAT, I am aware of the level of skill of one ordinarily skilled in the art of cardiovascular disease and kidney disease, and in particular, mechanisms of hypertension, hereto; AND being thus duly qualified declare as follows:

1. Much of my research has focused on animal models of hypertension, including studies in the SHR rat¹⁻⁴. This is a hereditary model of hypertension that we and others have shown is mediated by oxidative stress. While I am aware that xanthine oxidase inhibitors have been reported to lower blood pressure transiently in this model⁵ (also the Maeda Patent U.S. Patent 5,747,495), none of those studies suggested that this was due to uric acid, but rather asserted that this was due to oxidative stress. In fact, uric acid is considered an antioxidant and in some recent pilot studies we can show that it can lower blood pressure in these animals. Furthermore, most of the studies documenting oxidative stress in models of hypertension have focused on the role of NADPH oxidase, either produced by vascular cells^{6,7} or by the inflammatory cells themselves¹⁻⁴.

2. Dr Johnson was the first, as far as I am aware, to propose uric acid as a cause of hypertension, and the first to provide experimental evidence⁸. Specifically, he was also the first to suggest targeting uric acid levels with xanthine oxidase (x.o.) inhibitors as a means for improving blood pressure. While x.o. inhibitors could have some effects on blood pressure via their antioxidant effects, the observation that they failed to lower BP in longterm studies in the SHR ⁹⁻¹¹ taught away those skilled in the art from using them in

patients with hypertension. Furthermore, as mentioned above, I believe that most of the

oxidative stress in hypertension is due to a different system (NADPH oxidase) that is not

inhibited by xanthine oxidase inhibitors.

3. I therefore conclude that the concept of lowering uric acid as a means to control blood pressure was a novel idea for which Dr Johnson brought forth the first direct experimental⁸ as well as human¹² evidence. Studies in the SHR rat did not provide a reasonable expectation to those skilled in the art of successfully treating hypertension via administration of x.o. inhibitor, much less did they teach or suggest to those skilled in the

art to use x.o. inhibitors as a means to lower blood pressure by reducing uric acid levels.

4. The undersigned declares further that all statements made herein of his own knowledge are true and that all statements made on information in belief are believed to be true; and further that these statements were made with the knowledge that willful false statements in the like so made are punishable by fine or imprisonment, or both, under \$1001 of title 18 of the U.S.C. and that such willful false statements made jeopardize the

validity of the application or of any patent issuing thereon.

Further declarant sayeth naught.

Bernardo Rodríguez-Iturbe

October 24, 2007

Date

After freey

- **1.** Rodriguez-Iturbe B, Ferrebuz A, Vanegas V, Quiroz Y, Mezzano S, Vaziri ND. Early and sustained inhibition of nuclear factor-kappaB prevents hypertension in spontaneously hypertensive rats. *J Pharmacol Exp Ther* 2005;315:51-7.
- **2.** Rodriguez-Iturbe B, Quiroz Y, Nava M, Bonet L, Chavez M, Herrera-Acosta J, Johnson RJ, Pons HA. Reduction of renal immune cell infiltration results in blood pressure control in genetically hypertensive rats. *Am J Physiol Renal Physiol* 2002;282:F191-201.
- **3.** Nava M, Quiroz Y, Vaziri N, Rodriguez-Iturbe B. Melatonin reduces renal interstitial inflammation and improves hypertension in spontaneously hypertensive rats. *Am J Physiol Renal Physiol* 2003;284:F447-54.
- **4.** Rodriguez-Iturbe B, Zhan CD, Quiroz Y, Sindhu RK, Vaziri ND. Antioxidant-rich diet relieves hypertension and reduces renal immune infiltration in spontaneously hypertensive rats. *Hypertension* 2003;41:341-6.
- **5.** Miyamoto Y, Akaike T, Yoshida M, Goto S, Horie H, Maeda H. Potentiation of nitric oxide-mediated vasorelaxation by xanthine oxidase inhibitors. *Proc Soc Exp Biol Med* 1996;211:366-73.
- **6.** Rajagopalan S, Kurz S, Munzel T, Tarpey M, Freeman BA, Griendling KK, Harrison DG. Angiotensin II-mediated hypertension in the rat increases vascular superoxide production via membrane NADH/NADPH oxidase activation. Contribution to alterations of vasomotor tone. *J Clin Invest* 1996;97:1916-23.
- 7. Harrison DG, Cai H, Landmesser U, Griendling KK. Interactions of angiotensin II with NAD(P)H oxidase, oxidant stress and cardiovascular disease. *J Renin Angiotensin Aldosterone Syst* 2003;4:51-61.
- **8.** Mazzali M, Hughes J, Kim YG, Jefferson JA, Kang DH, Gordon KL, Lan HY, Kivlighn S, Johnson RJ. Elevated uric acid increases blood pressure in the rat by a novel crystal-independent mechanism. *Hypertension* 2001;38:1101-6.
- **9.** Laakso J, Mervaala E, Himberg JJ, Teravainen TL, Karppanen H, Vapaatalo H, Lapatto R. Increased kidney xanthine oxidoreductase activity in salt-induced experimental hypertension. *Hypertension* 1998;32:902-6.
- **10.** Maenishi O, Ito H, Suzuki T. Acceleration of hypertensive cerebral injury by the inhibition of xanthine-xanthine oxidase system in stroke-prone spontaneously hypertensive rats. *Clin Exp Hypertens* 1997;19:461-77.
- **11.** Trachtman H, Valderrama E, Futterweit S. Nephrotoxicity of allopurinol is enhanced in experimental hypertension. *Hypertension* 1991;17:194-202.
- **12.** Feig DI, Nakagawa T, Karumanchi SA, Oliver WJ, Kang DH, Finch J, Johnson RJ. Hypothesis: Uric acid, nephron number, and the pathogenesis of essential hypertension. *Kidney Int* 2004;66:281-7.
- **13.** Ward HJ. Uric acid as an independent risk factor in the treatment of hypertension. *Lancet* 1998;352:670-1.
- **14.** Culleton BF, Larson MG, Kannel WB, Levy D. Serum uric acid and risk for cardiovascular disease and death: the Framingham Heart Study. *Ann Intern Med* 1999;131:7-13.

Bernardo Rodríguez.Iturbe CURRICULUM-VITAE

Personal data:

NAME: BERNARDO RODRIGUEZ-ITURBE

CI. 1.686.802

Birth plce and date MARACAIBO, VENEZUELA, January 4, 1939

NACIONALITY (Country) VENEZUELA

Wife: MARIA TERESA ESPINOSA

Children: MARIA TERESA, BERNARDO, PAULINA, DANIELA

Prsesent Academic Appointments

Profesor of Medicine Facultad de Medicina, Universidad del Zulia,

Maracaibo, Venezuela.

Medical Director, Renal

Transplant Team Hospital Universitario de Maracaibo, Maracaibo,

Venezuela

Chief, Nephrology Service Hospital Universitario de Maracaibo, Venezuela

Director, Postgrado Neprohology

Residency Program Universidad del Zulia, Facultad de Medicina, Div. de

Estudios de Postgrado y Hospital Universitario de

Maracaibo.

Director Centro de Investigaciones Biomédicas, Instituto Venezolano de

Investigaciones Cientificas - Zulia (IVIV-Zulia

Academic Degrees

M.D. Facultad de Medicina, Universidad del

Zulia, Maracaibo, Venezuela 1961

Master University of Pennsylvania

Medical Sciences Graduate School of Medicine,

Philadelphia, Pa. 1967

Doctor Facultad de Medicina,

Ciencias Médicas Universidad del Zulia

Maracaibo, Venezuela 1970

Past Academic Appointments	INSTITUTION	<u>DATE</u>
Coordinador-Director Postgraduate Studies Office Postgrado	Universidad del Zulia, Facultad de Medicina	1975-1978
School of Medicine Council Member	Universidad del Zulia, Escuela de Medicina	1974-1976 1978-1980
Faculty Council Member	Universidad del Zulia, Facultad de Medicina	1976-1979
Director	Instituto de Investigaciones, Biomédicas (INBIOMED)	1983-
Chief of Medical Clinic	Universidad del Zulia, Facultad de Medicina	1985-1987
Chairman of Department Medicine	Universidad del Zulia, Facultad de Medicina	1978-1980 1988-1989
Past Hospital Appointments		
Medical Director	Intensive Care Unit Hospital Universitario de Maracaibo	1968-1971
Director	Equipo de Trasplantes Renales Hospital Universitario de Maracaibo	1968-
Medical Director	Hospital Universitario de Maracaibo	1981-1983
Médico Jefe II	Servicio de Nefrología Hospital Universitario de Maracaibo	1983-
Other Institutions	INSTITUTION	<u>DATE</u>
Consultor	Organización Mundial de la Salud	Feb. 1972
Member Scientific Committe	Zulia Medical College	1969-1971
	Universidad del Zulia	1976-1979

Council Member	Consejo Nacional de Diálisis y Trasplante	1977-
Member, Committee for Clinical Medical Sciences . (Chairman)	Consejo Nacional Invest. Científicas y Tecnológicas (CONICIT)	1973-1978 1976-1978
Chairman, Task Force	Prersidential Task force for the Evaluation and Promotion of Science and Technology in Zulia region	1979
Council member, Postgraduate Studies y Tecr	Consejo Nacional Invest. Científicas nológicas (CONICIT) 1983	-1984
Council member,	Sistema de Promoción del Investigador (PPI)	1992-1996
Member	Academia de Ciencias de América Latina (ACAL)	1993-
Presidente FUNDASALUD	Gobernación, Estado Zulia	1994-1998
Delegado Principal CONICIT	Consejo Nacional de Universidades	1994-
Councilman	International Society of Nephrology	1995-2001
Member, Commission for the Global Advancement of Nephrology (COMGAN)	International Society of Nephrology Chariman, Latin American Subcommitte	1995-2005
President	Panamerican Society of Dialysis and Transplantation	1995-1997
Co-Director	Comité de Curriculum y Conocimientos Básicos de Nefrología, Sociedad Latinoamericana de Nefrología e Hipertensión	1996-1999
Miembro, Consejo de Apelaciones , Sistema de Promoción al Investigador (CONICIT)		2001-
Member, Prize and Awards Comission Latinamerican Society of Nephrology and Hypertension (SLANH) 2003		
Member Fellowship Comité, Internacional Society of Nephrology		2004-2005

Awards during Medical School

Award for the highest academic score In Medical Studies	Escuela de Medicina Universidad del Zulia	1957-1958 1958-1959
Joaquín Esteva Parra Awards	Facultad de Medicina Universidad del Zulia	1959
Brancisco Eugenio Bustamante Award	Universidad del Zulia	1960
Graduation SUMMA CUM LAUDE	Facultad de Medicina, Universidad del Zulia,	1961
Postgradute Scholarships Short-Term Scholarship	American College of Physicians	1973
Awards and Professional Recognitions		
Prize CONICIT to the best Research paper Cier	Consejo Nacional de Ciencia y ncia de la Salud Tecnología (CONICIT)	1977
Order Jesús Enrique Lossada	Universidad del Zulia	1986
Prize Scientific Merit	FUNDACITE-ZULIA	1986
Order Andrés Bello, For contributions to Science	República de Venezuela	1990
Research Award Francisco Eugenio Bustamante	Universidad del Zulia	1993
Prize for Clinical Investigation	Sociedad Venezolana de Nefrología	1993
Laureate Award	American College of Physicians	1995
PREMIO NACIONAL DE CIENCIA	Venezuela (CONICIT)	1998
PREMIO "LORENZO MENDOZA FLEUR	Y" Fundación Polar (Venezuela)	1999
PREMIO "VÍCTOR RAÚL MIATELLO" Sociedad Latinoamericana de Nefrología		1999
INTERNATIONA MEDAL,	National Kidney Foundation	2004

Luis Hernando	International	Prize
in Nephrology		

Fundación Iñigo Alvarez de Toledo

2007

Other Recognitions

Investigador Nivel IV Sistema de Promoción al Investigador (PPI)	1990
Doctorado Honoris Causa Universidad del Zulia	2001
International Fellow of The Council for High Blood Pressure Research	2002
Profesor Honoris Causa, Sociedad Mexicana de Hipertension Arterial	2006
Miembro de Honor, Sociedad Española de Nefrología	2006
Jose Strauss Award, Pediatric Nephrology Seminar, Miller School of Medicine, Miami University	2007

Editorial Board of Scientific Journals

Asesor Científico	INVESTIGACION CLINICA Maracaibo, Venezuela	1973-1982
Comité Editorial	REVISTA DE LA ACADEMIA DE MEDICINA, Maracaibo, Venezuela	1988-
Editorial Board	CLINICAL NEPHROLOGY München, Diesenhofen, FRG	1975-1995
Comité Editorial	CIENCIA, Facultad Experimental de Ciencias Maracaibo, Venezuela	1993-
Comité Editorial	NEFROLOGIA LATINOAMERICANA (Soc. Latinoamericana de Nefrologia)	1993-
Editor	KIDNEY INTERNATIONAL SPANISH AND PORTUGUESE	2005-
Editorial Board	AMERICAN JOURNAL OF KIDNEY DISEASES	2004- 2006
Advisory Board	NATURE, Clinical Practice Nephrology	2005
Editorial Board	Clinical Journal of the	

	American Society of Nephrology (CJASN)		2005-2006
Editorial Board	Current Opinion Nephrology and Hypertension		2007-2009
Medical and Postgrad	<u>luate studies</u>	INSTITUCION	<u>FECHA</u>
Estudios de Medicina		FacItad de Medicina Uiversidad del Zulia Maracaibo, Venezuela	1955-1961
Estudios de Postgrado		Postgraduate School of Medicine, University of Pennsylvania, School of Medicine, Philadelphia, PA.	1961-1962
Residencia Medicina Interna		Graduate Hospital, University of Pennsylvania, Philadelphia, PA.	1962-1965
Fellowship, Nefrologia	ı	Graduate Hospital and University of Penssylvania, Medical Center, Philadelphia, PA.	1964-1966

Medical Societies

Available upon request

Invited Conferences, Plenary lectures, Visiting Professorships Available upon request.

Abstracts and presentations. Available upon request.

PUBLICATIONS IN SCIENTIFIC JOURNALS

- 1. HENDERSON LW, RODRIGUEZ ITURBE B, BLUEMLE LW. Factors influencing b lood pH changes during extracorporeal hemodialysis in patients with chronic renal failure. *Trans Amer Soc Artif Intern Organs* 12:193-199, 1966.
- 2. RODRIGUEZ-ITURBE B, VERA G, RIVERA H, SERRANO H, SHAW HJ, GARCIA R. Homotrasplante renal. *Inves. Clin* 24:9-25, 1967.
- 3. RODRIGUEZ-ITURBE B. Evolución postoperatoria de los trasplantes renales. *Gaceta Médica de Caracas* 1:31-42, 1969.
- 4. RODRIGUEZ-ITURBE B, GARCIA R, MEDINA A, RUBIO L, SERRANO H. Curso postoperatorio del trasplante renal. *Acta Méd Venez* 16:99-103, 1969.
- 5. SERRANO H, RODRIGUEZ-ITURBE B. Estudio inmunológicos como ayuda en la detección del rechazo en el homoinjerto renal. *Invest Clin* 29:9-27, 1969.
- 6. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L. Diálisis crónica. Experiencia en el Hospital Universitario de Maracaibo. *Invest Clin* 31:7-43, 1969.
- 7. SERRANO H, RODRIGUEZ-ITURBE B, GARCIA R, CUENCA L. Complemento sérico en la glomerulonefritis aguda. *Invest Clin* 33:19-29, 1970.
- 8. RIVERA H, ALONZO A, RODRIGUEZ-ITURBE B, MATA H, GONZALEZ F, MOLERO M, BUCOBO E. Aspectos técnicos vasculares de los trasplantes renales. *Bol Soc Ven Cirugía* 24:717-722, 1970.
- 9. RODRIGUEZ-ITURBE B. Acute yellow phosphorus poisoning (Editorial). *New Engl J Med* 284:157, 1971.
- 10. RODRIGUEZ-ITURBE B, SERRANO H, GARCIA R, GALLEGOS B. Reliability of changes of serum complement, C3 and immunoglobulins during acute rejection of renal allografts. *Transplantation* 12: 405-407, 1971.
- 11. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L, ZABALA J, MOROS G, TORRES R. Acute glomerulonephritis in the Guillain-Barré-Strohl syndrome. Report of 9 cases. *Ann intern Med* 78:391-395, 1973.
- 12. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L, HENDERSON LW. Quantitation of hydrogen ion removal by extracorporeal hemodialysis in patiens with chronic renal failure. *Clin Nephrol* 2:238-244, 1974.
- 13. GARCIA R, RODRIGUEZ-ITURBE B, RUBIO L. Hemodiálisis en el hogar. Experiencia en el Hospital Universitario de Maracaibo. *Invest Clin* 16:143-163, 1975.
- 14. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L, TRESER G, LANGE K. Epidemic glomerulonephritis in Maracaibo. Evidence for progression to chronicity. *Clin Nephrol* 5:107-206, 1976.

- 15. LAYRISSE Z, PULIDO DE RODRIGUEZ M, RODRIGUEZ-ITURBE B, GARCIA E, STOILOW Z, SALAS G. Genetics of the HLA-A system in Venezuelan heterogenous population. *Vox Sang* 31:37-47, 1976.
- 16. GARCIA R, RODRIGUEZ-ITURBE B, RUBIO L, FERRER J. Hemodiálisis por doble cateterización de vena femoral. *Invest Clin* 17:122-127, 1976.
- 17. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L, SERRANO H. Immunohistologic findings in the lung in systemic lupus erythematosus. *Arch Pathol Lab Med* 101: 342-344, 1977.
- 18. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L. Glomeruonephritis post estreptocóccica. Aspectos controveriales de investigación reciente. La enfermedad en Venezuela. *Acta Cientif Venez* 28:245-248, 1977.
- 19. RODRIGUEZ-ITURB E B, GARCIA R, RUBIO L, LAYRISSE Z, MORALES E. Trasplante renal y diálisis crónica: estudios de sobrevida e histocompatiblidad. Incidencia de tuberculosis. *Invest Clin* 18:136-145, 1977.
- 20. RODRIGUEZ-ITURBE B. Algunos aspectos de la regulación renal de la tensión arterial. Revisión. *Invest Clin* 19(4): 156-181, 1978.
- 21. RODRIGUEZ-ITURBE B. Venezuela: Health Care and Medical Education. *Ann Intern Med* 88:125-126, 1978.
- 22. RODRIGUEZ-ITURBE B. Comentarios sobre la situación curricular de los estudios médicos en la Universidad del Zulia. *Acta Cientif Venez* 29:143-146, 1978.
- 23. RODRIGUEZ-ITURBE B, GARCIA R, WEBSTER GD, RUBIO L. Diálisis peritoneal y diuresis forzada en el tratamiento de la intoxicación por Glutethimida. *Invest. Clin* 19:87-101, 1978.
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- 25. McINTOSH RM, GARCIA R, RUBIO L, RABIDEAU D, ALLEN JE, CARR RI, RODRIGUEZ-ITURBE B. Evidence for an autologous immune complex pathogenic mechanism in acute poststreptococcal glomerulonephritis. *Kidney Int* 14:501-510, 1978
- 26. McINTOSH RM, RABIDEAU D, ALLEN JE, GARCIA R, RUBIO L, CARR RI, RODRIGUEZ-ITURBE B. Acute poststreptococcal glomerulonephritis in Maracaibo II. Studies on the incidence, nature and significance of circulating antiimmunoglobulins. *Ann Rheum Dis* 38:257-261, 1979.
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- 28. RODRIGUEZ-ITURBE B, CASTILLO L, VALBUENA R, CUENCA L. Acute poststreptococcal glomerulonephritis. A review of recent developments. *Pediatrician* 8:307-324, 1979.
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- 30. RODRIGUEZ-ITURBE B, CARR RI, GARCIA R, RABIDEAU D, RUBIO L, McINTOSH RM. Circulating immune complexes and serum immunoglobulins in acute poststreptococcal glomerulonephritis- Evidence for circulating immune complex pathogenesis. *Clin Nephrol* 13:1-5, 1980.
- 31. RODRIGUEZ-ITURBE B, RABIDEAU D, GARCIA R, RUBIO L, McINTOSH RM, Characterization of the glomerular antibody in acute poststreptococcal glomerulonephritis. *Ann Intern Med* 92:478-481, 1980.
- 32. RODRIGUEZ-ITURBE B, BAGGIO B, COLINA-CHOURIO J, FAVARO S, GARCIA R, SUSSANA F, CASTILLO L, BORSATTI A. Studies on the renin-aldosterone system in the acute nephritic syndrome. *Kidney Int* 19:445-453, 1981.
- 33. RODRIGUEZ-ITURBE B, RUBIO L, GARCIA R. Attack rate of poststreptococcal nephritis in families. A prospective study. *Lancet* 1: 401-403, 1981.
- 34. GARCIA R, RUBIO L, RODRIGUEZ-ITURBE B. Long-term prognosis of epidemic poststreptococcal glomerulonephritis in Maracaibo: follow-up studies 11-12 years after the acute episode. *Clin Nephrol* 15: 291-298, 1981.
- 35. RODRIGUEZ-ITURBE B, KATIYAR VN, COELLO J. Neuraminidase activity and free sialic acid levels in the serum of patients with acute poststreptococcal glomerulonephritis. *New Engl J. Med* 304:1506-1510, 1981.
- 36. RODRIGUEZ-ITURBE B, SILVA BEAUPERHUY V, PARRA G, RUBIO L, GARCIA R. Skin window immune response to normal human IgG in patients with rheumatoid arthritis and acute poststreptococcal glomerulonephritis. *Amer J Clin Pathol* 76: 270-275, 1981.
- 37. RODRIGUEZ-ITURBE B, MORENO-FUENMAYOR H, RUBIO L, GARCIA R, LAYRISSE Z. Mendelian recessive ratios in acute poststreptococcal glomerulonephritis. *Experentia* 38: 918-920, 1982.
- 38. LAYRISSE A, RODRIGUEZ-ITURBE B, GARCIA R, RODRIGUEA A, TIWARI J. Family studies of HLA system in acute poststreptococcal glomerulonephritis. *Human Immunol* 7:177-185, 1983.
- 39. RIVERA S, BELLOSO H, RINCON R, RODRIGUEZ-ITURBE B. Ensayo de inducción de tolerancia específica a injertos de piel mediante trasplante de córnea en ratones. *Invest Clin* 24: 99-107, 1983.
- 40. COLINA -CHOURIO J., RODRIGUEZ-ITURBE B, BAGGIO B, GARCIA R, BORSATTI A. Urinary excretion of prostaglandins (PGE2 and PGF2a) and kallidrein in acute glomerulonephritis. *Clin Nephrol* 20:217-224, 1983.
- 41. VOGT A, BATSFORD SR. RODRIGUEZ-ITURBE B, GARCIA R, Cationic antigens in poststreptococcal glomerulonephritis. *Clin Nephrol* 20: 271-279, 1983.
- 42. HENRIQUEZ-LA ROCHE C, RODRIGUEZ-ITURBE B. Infección del tracto urinario. Tópicos de importancia y tratamiento. Revisión. *Invest Clin* 25: 103-118, 1984.
- 43. COLINA-CHOURIO J, RODRIGUEZ-ITURBE B. Usos clínicos de los inhibidores de la enzima de conversión. *Invest Clin* 25: 33-47, 1984.

- 44. MOSQUERA J, RODRIGUEZ-ITURBE B. Extracellular neuraminidase production of streptococci associated with acute nephritis. *Clin Nephrol* 21: 21-28, 1984.
- 45. RODRIGUEZ-ITURBE B. Epidemic poststreptococcal glomerulonephritis (Nephrology Forum). *Kidney Int* 25: 129-136, 1984.
- 46. RODRIGUEZ-ITURBE B. La historia natural de la nefritis postestreptocóccica. N Arch Fac Med 42: 457-460, 1984.
- 47. PARRA G, PLATT JL, FLAK RJ, RODRIGUEZ-ITURBE B, MICHAEL AF. Cell populations and membrane attack complex in glomeruli of patients with poststreptococcal glomerulonephritis: identification using monoclonal antibodies by indirect immunofluorescence. *Clin Immunol Immunopathol* 33: 324-332, 1984.
- 48. GARCIA RAMIREZ R, RUBIO L, RODRIGUEZ-ITURBE B. Indicaciones y contraindicaciones del trasplante renal.. Revisión. *Invest Clin* 25: 213-237, 1984.
- 49. MOSQUERA JA, KATIYAR VN, COELLO J, RODRIGUEZ-ITURBE B. Extracellular neuraminidase production of streptococci isolated from patients with glomerulonephritis. *J Infect Dis* 151:259-263, 1985.
- 50. RODRIGUEZ-ITURBE B, GARCIA R, RUBIO L, CUENCA L. . Características clínicas y epidemilógicas de la glomerulonefritis postestreptocóccica en la región zuliana. *Invest Cli*n 26 (3): 191-211, 1985.
- 51. WONG A, GARCIA R, HERIQUEZ C, RODRIGUEZ-ITURBE B. Salmonella osteomyelitis in a renal trasplant recipient. *Brief Report. Invest Clin* 26 (4): 231-234, 1985.
- 52. RODRIGUEZ-ITURBE B, HERRERA J, GARCIA R. Response to acute protein load in kidney donors and in apparently normal postacute glomerulonphritis patients: evidence for glomerular hyperfiltration. *Lancet* 2: 461-464. 1985.
- 53. RODRIGUEZ-ITURBE B. The functional reserve capacity of the kidney. *Int J Artf Org* 9 (2): 81-84, 1986.
- 54. MOSQUERA J, RODRIGUEZ-ITURBE B. Glomerular binding sites for peanut agglutinin in acute poststretococcal glomerulonephritis. *Clin Nephrol* 26 (5): 227-234, 1986.
- 55. HERRERA J, GARCIA R, RODRIGUEZ-ITURBE B. Parasitosis in the immunosuppressed host. Review. *Invest Clin* 28 (1): 47-59, 1987.
- 56. COOK GA, RODRIGUEZ H, SILVA H, RODRIGUEZ-ITURBE B, BOHORQUEZ H. Adult respiratory distress secondary to strongyloidiasis. *Chest* 92: 1115.116, 1987.
- 57. HERRERA J, GARCIA R, RUBIO L, HENRIQUEZ C, RODRIGUEZ-ITURBE B. Sobrevida actuarial en trasplante renal: Hospital Universitario de Maracaibo (1967-1986). *Invest Clin* 28(3): 133-142, 1987.

- 58. NAVARRO JA, GARCIA R, RUBIO D, RODRIGUEZ-ITURBE B, VIRLA J, MENDEZ G, ROMERO R. Correlación de los niveles de aluminio en agua, sangre completa y dializado de pacientes en hemodiálisis crónica. *Invest Clin* 28(3) 153-163, 1987.
- 59. RODRIGUEZ-ITURBE B, HERRERA J, GARCIA R. Relationship between glomerular filtrtion rate and renal blood flow at different levelvs of protein-induce hyperfiltarion in man. *Clin Sci* 74:11-15, 1988.
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